

RECEIVED

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<110> MACK, DAVID GISH, KURT

<120> NOVEL METHODS OF DIAGNOSING AND TREATING BREAST CANCER, COMPOSITIONS, AND METHODS OF SCREENING FOR BREAST CANCER MODULATORS

- <130> A-67860-3/DJB/JJD
- <140> US 09/525,361
- <141> 2000-03-15
- <150> US 09/268,865
- <151> 1999-03-15
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- <151> 1999-11-29
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- <150> US 09/520,478
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- <150> US 09/440,676
- <151> 1999-11-16
- <150> US 09/440,677
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45

40

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- Leu Thr Leu Asp Ile Ala Ser Ile Ser Ser Ile Ser Glu Ser Asn Met 65 70 75 80
- Asp Tyr Thr Ala Thr Ile Tyr Leu Arg Gln Arg Trp Met Asp Gln Arg 85 90 95
- Leu Val Phe Glu Gly Asn Lys Ser Phe Thr Leu Asp Ala Arg Leu Val
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- Glu Phe Leu Trp Val Pro Asp Thr Tyr Ile Val Glu Ser Lys Lys Ser 115 120 125
- Phe Leu His Glu Val Thr Val Gly Asn Arg Leu Ile Arg Leu Phe Ser 130 135 140
- Asn Gly Thr Val Leu Tyr Ala Leu Arg Ile Thr Thr Thr Val Ala Cys 145 150 155 160
- Asn Met Asp Leu Ser Lys Tyr Pro Met Asp Thr Gln Thr Cys Lys Leu 165 170 175
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- Leu Arg Gly Asn Asp Ser Val Arg Gly Leu Glu His Leu Arg Leu Ala 195 200 205
- Gln Tyr Thr Ile Glu Arg Tyr Phe Thr Leu Val Thr Arg Ser Gln Gln 210 215 220
- Glu Thr Gly Asn Tyr Thr Arg Leu Val Leu Gln Phe Glu Leu Arg Arg 225 230 235 240
- Asn Val Leu Tyr Phe Ile Leu Glu Thr Tyr Val Pro Ser Thr Phe Leu 245 250 255
- Val Val Leu Ser Trp Val Ser Phe Trp Ile Ser Leu Asp Ser Val Pro 260 265 270
- Ala Arg Thr Cys Ile Gly Val Thr Thr Val Leu Ser Met Thr Thr Leu 275 280 285
- Met Ile Gly Ser Arg Thr Ser Leu Pro Asn Thr Asn Cys Phe Ile Lys 290 295 300
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Leu Leu Glu Tyr Ala Val Ala His Tyr Ser Ser Leu Gln Gln Met Ala 325 330 335

Ala Lys Asp Arg Gly Thr Thr Lys Glu Val Glu Glu Val Ser Ile Thr 340 345 350

Asn Ile Ile Asn Ser Ser Ile Ser Ser Phe Lys Arg Lys Ile Ser Phe 355 360 365

Ala Ser Ile Glu Ile Ser Ser Asp Asn Val Asp Tyr Ser Asp Leu Thr 370 375 380

Met Lys Thr Ser Asp Lys Phe Lys Phe Val Phe Arg Glu Lys Met Gly 385 390 395

Arg Ile Val Asp Tyr Phe Thr Ile Gln Asn Pro Ser Asn Val Asp His
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- Ser Val Thr Glu Ser Pro Gln Arg Val Ile Ile Thr Glu Asp Asp Glu 50 55 60
- Asp Glu Thr Thr Val Glu Leu Glu Gly Gln Asp Glu Asn Gln Glu Gly 65 70 75 80
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- Tyr Asp Asp Glu Glu Phe Glu Gly Tyr Glu Asp Lys Pro Asp Thr Ser 100 105 110
- Ser Ser Lys Asn Lys Asp Pro Ile Thr Ile Val Asp Val Pro Ala His 115 120 125
- Leu Gln Asn Ser Trp Glu Ser Tyr Tyr Leu Glu Ile Leu Met Val Thr 130 135 140
- Gly Leu Leu Ala Tyr Ile Met Asn Tyr Ile Ile Gly Lys Asn Lys Asn 145 150 155 160
- Ser Arg Leu Ala Gln Ala Trp Phe Asn Thr His Arg Glu Leu Leu Glu 165 170 175
- Ser Asn Phe Thr Leu Val Gly Asp Asp Gly Thr Asn Lys Glu Ala Thr 180 185 190
- Ser Thr Gly Lys Leu Asn Gln Glu Asn Glu His Ile Tyr Asn Leu Trp 195 200 205
- Cys Ser Gly Arg Val Cys Cys Glu Gly Met Leu Ile Gln Leu Arg Phe 210 215 220
- Leu Lys Arg Gln Asp Leu Leu Asn Val Leu Ala Arg Met Met Arg Pro 225 230 235 240
- Val Ser Asp Gln Val Gln Ile Lys Val Thr Met Asn Asp Glu Asp Met 245 250 255
- Asp Thr Tyr Val Phe Ala Val Gly Thr Arg Lys Ala Leu Val Arg Leu 260 265 270

Gln Lys Glu Met Gln Asp Leu Ser Glu Phe Cys Ser Asp Lys Pro Lys 275 280 285

Ser Gly Ala Lys Tyr Gly Leu Pro Asp Ser Leu Ala Ile Leu Ser Glu 290 295 300

Met Gly Glu Val Thr Asp Gly Met Met Asp Thr Lys Met Val His Phe 305 310 315 320

Leu His Thr Tyr Ala Asp Lys Ile Glu Ser Val His Phe Ser Asp Gln 325 330 335

Phe Ser Gly Pro Lys Ile Met Gln Glu Glu Gly Gln Pro Leu Lys Leu 340 345 350

Pro Asp Thr Lys Arg Thr Leu Leu Phe Thr Phe Asn Val Pro Gly Ser 355 360 365

Gly Asn Thr Tyr Pro Lys Asp Met Glu Ala Leu Leu Pro Leu Met Asn 370 375 380

Met Val Ile Tyr Ser Ile Asp Lys Ala Lys Lys Phe Arg Leu Asn Arg 385 390 395 400

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90

Phe Ala Leu Cys Gly Pro Gln Met Leu Val Phe Leu Arg Val Ile Gly

85

100 105 110

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Tyr Pro Val Lys Tyr Thr Gln Thr Phe Thr Leu His Ala Asn Pro Ala 130 135 140

Val Thr Tyr Ile Tyr Asn Trp Ala Tyr Gly Phe Gly Trp Ala Ala Thr 145 150 155 160

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Val Asn Tyr Ile Tyr Asn Trp Ala Tyr Gly Phe Gly Trp Ala Ala Thr 145 . 150 155 160

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Thr Gln Thr Phe Arg Leu His Asp Asn Pro Ala Val Asn Tyr Ile Tyr 65 70 75 80

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Lys Gln Lys Arg Thr Cys Tyr Leu Gly Ser Lys Thr Leu Phe Tyr Arg
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60

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Thr Met Tyr Phe Phe Phe Gly Leu Leu Gly Val Ala Asp Ile Leu Cys 115 120 125

Phe Thr Ile Ser Ser Leu Pro Val Ser Leu Thr Lys Leu Met Leu Ser 130 135 140

Asn Ala Leu Phe Val Glu Ala Phe Ile Phe Tyr Asn His Thr His Gly 145 150 155 160

Arg Glu Met Leu Asp Ile Phe Val His Gln Leu Leu Val Leu Val Val 165 170 175

Phe Leu Thr Gly Leu Val Ala Phe Leu Glu Phe Leu Val Arg Asn Asn 180 185 190

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Trp Phe Phe Gln Ile Gly Phe Val Leu Tyr Pro Pro Ser Gly Gly Pro 210 215 220

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Phe Cys Trp His Tyr Ala Val Thr Ile Val Ile Val Gly Met Asn Tyr 245 250 255

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Ser Asp Ala Ala Glu Leu Asn His Lys Glu Glu His Ser Leu His Val 50 55 60

Gln Asp Pro Ser Ser Ser Ser Lys Lys Asp Leu Lys Ser Ala Val Leu 65 70 75 80

Ser Glu Lys Ala Gly Phe Asn Tyr Glu Ser Pro Ser Lys Gly Gly Asn 85 90 95

Phe Pro Ser Phe Pro His Asp Glu Val Thr Asp Arg Asn Met Leu Ala
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Gly Asp Ser Leu Glu Thr Lys Glu Asp Gln Lys Met Ser Pro Lys Ala 145 150 155 160

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Gly Gly Val Arg Leu Asn Lys Ser Lys Thr Asp Leu Leu Val Asn Asp 195 200 205

Asn Pro Asp Pro Ala Pro Leu Ser Pro Glu Leu Gln Asp Phe Lys Cys 210 215 220

Asn Ile Cys Gly Tyr Gly Tyr Gly Asn Asp Pro Thr Asp Leu Ile 225 230 235 240

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- Asp Ala Glu Leu Asp Ser Lys Ile Leu Ala Leu His Asn Met Val Gln 260 265 270
- Phe Ser His Ser Lys Asp Phe Gln Lys Val Asn Arg Ser Val Phe Ser 275 280 285
- Gly Val Leu Gln Asp Ile Asn Ser Ser Arg Pro Val Leu Leu Asn Gly 290 295 300
- Thr Tyr Asp Val Gln Val Thr Ser Gly Gly Thr Phe Ile Gly Ile Gly 305 310 315 320
- Arg Lys Thr Pro Asp Cys Gln Gly Asn Thr Lys Tyr Phe Arg Cys Lys 325 330 335
- Phe Cys Asn Phe Thr Tyr Met Gly Asn Ser Ser Thr Glu Leu Glu Gln 340 345 350
- His Phe Leu Gln Thr His Pro Asn Lys Ile Lys Ala Ser Leu Pro Ser 355 360 365
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- Ile Thr Val Lys Ala Gly Asp Asp Thr Pro Val Gly Tyr Ser Val Pro
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- Leu Lys Leu Leu Glu His Tyr Gly Lys Gln His Gly Ala Val Gln Ser 450 455 460
- Gly Gly Leu Asn Pro Glu Leu Asn Asp Lys Leu Ser Arg Gly Ser Val 465 470 475 480
- Ile Asn Gln Asn Asp Leu Ala Lys Ser Ser Glu Gly Glu Thr Met Thr
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- Lys Thr Asp Lys Ser Ser Ser Gly Ala Lys Lys Lys Asp Phe Ser Ser 500 505 510

- Lys Gly Ala Glu Asp Asn Met Val Thr Ser Tyr Asn Cys Gln Phe Cys 515 520 525
- Asp Phe Arg Tyr Ser Lys Ser His Gly Pro Asp Val Ile Val Val Gly 530 535 540
- Pro Leu Leu Arg His Tyr Gln Gln Leu His Asn Ile His Lys Cys Thr 545 550 555 560
- Ile Lys His Cys Pro Phe Cys Pro Arg Gly Leu Cys Ser Pro Glu Lys
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- His Leu Gly Glu Ile Thr Tyr Pro Phe Ala Cys Arg Lys Ser Asn Cys 580 585 590
- Ser His Cys Ala Leu Leu Leu His Leu Ser Pro Gly Ala Ala Gly 595 600 605
- Ser Ser Arg Val Lys His Gln Cys His Gln Cys Ser Phe Thr Thr Pro 610 615 620
- Asp Val Asp Val Leu Leu Phe His Tyr Glu Ser Val His Glu Ser Gln 625 630 635 640
- Ala Ser Asp Val Lys Gln Glu Ala Asn His Leu Gln Gly Ser Asp Gly 645 650 655
- Gln Gln Ser Val Lys Glu Ser Lys Glu His Ser Cys Thr Lys Cys Asp
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- Asp Ser Lys Met Gly Glu Pro Val Ser Glu Ser Val Val Lys Arg Glu 755 760 765
- Lys Leu Glu Glu Lys Asp Gly Leu Lys Glu Lys Val Trp Thr Glu Ser 770 780

- Ser Ser Asp Asp Leu Arg Asn Val Thr Trp Arg Gly Ala Asp Ile Leu 785 790 795 800
- Arg Gly Ser Pro Ser Tyr Thr Gln Ala Ser Leu Gly Leu Leu Thr Pro 805 810 815
- Val Ser Gly Thr Gln Glu Gln Thr Lys Thr Leu Arg Asp Ser Pro Asn 820 825 830
- Val Glu Ala Ala His Leu Ala Arg Pro Ile Tyr Gly Leu Ala Val Glu 835 840 845
- Thr Lys Gly Phe Leu Gln Gly Ala Pro Ala Gly Glu Lys Ser Gly 850 855 860
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- Gly Gly Tyr Val Cys Asn Ala Cys Gly Leu Tyr Gln Lys Leu His Ser 915 920 925
- Thr Pro Arg Pro Leu Asn Ile Ile Lys Gln Asn Asn Gly Glu Gln Ile 930 935 940
- Ile Arg Arg Arg Thr Arg Lys Arg Leu Asn Pro Glu Ala Leu Gln Ala 945 950 955 960
- Glu Gln Leu Asn Lys Gln Gln Arg Gly Ser Asn Glu Glu Gln Val Asn 965 970 975
- Gly Ser Pro Leu Glu Arg Arg Ser Glu Asp His Leu Thr Glu Ser His 980 985 990
- Gln Arg Glu Ile Pro Leu Pro Ser Leu Ser Lys Tyr Glu Ala Gln Gly 995 1000 1005
- Ser Leu Thr Lys Ser His Ser Ala Gln Gln Pro Val Leu Val Ser Gln 1010 1015 1020
- Thr Leu Asp Ile His Lys Arg Met Gln Pro Leu His Ile Gln Ile Lys 1025 1030 1035 1040
- Ser Pro Gln Glu Ser Thr Gly Asp Pro Gly Asn Ser Ser Ser Val Ser 1045 1050 1055

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Arg Pro Ala Lys His Pro Asn Tyr Ser Pro Pro Gly Ser Pro Ile Glu 1075 1080 1085

Lys Tyr Gln Tyr Pro Leu Phe Gly Leu Pro Phe Val His Asn Asp Phe 1090 1095 1100

Gln Ser Glu Ala Asp Trp Leu Arg Phe Trp Ser Lys Tyr Lys Leu Ser 1105 1110 1115 1120

Val Pro Gly Asn Pro His Tyr Leu Ser His Val Pro Gly Leu Pro Asn 1125 1130 1135

Pro Cys Gln Asn Tyr Val Pro Tyr Pro Thr Phe Asn Leu Pro Pro His 1140 1145 1150

Phe Ser Ala Val Gly Ser Asp Asn Asp Ile Pro Leu Asp Leu Ala Ile 1155 1160 1165

Lys His Ser Arg Pro Gly Pro Thr Ala Asn Gly Ala Ser Lys Glu Lys 1170 1175 1180

Thr Lys Ala Pro Pro Asn Val Lys Asn Glu Gly Pro Leu Asn Val Val 1185 1190 1195 1200

Lys Thr Glu Lys Val Asp Arg Ser Thr Gln Asp Glu Leu Ser Thr Lys 1205 1210 1215

Cys Val His Cys Gly Ile Val Phe Leu Asp Glu Val Met Tyr Ala Leu 1220 1225 1230

His Met Ser Cys His Gly Asp Ser Gly Pro Phe Gln Cys Ser Ile Cys 1235 1240 1245

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- Gly Lys Tyr Val His Asn His Asn Val Tyr Thr Asn Asn Glu Asn Cys
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- Ser Ser Pro Ser Trp Gln Ala Met His Glu Pro Arg Thr Phe Ala Val 115 120 125
- Tyr Leu Asn Asn Thr Gly Tyr Arg Thr Ala Phe Phe Gly Lys Tyr Leu 130 135 140
- Gly Leu Ile Lys Asn Ser Arg Phe Tyr Asn Tyr Thr Val Cys Arg Asn 165 170 175
- Gly Ile Lys Glu Lys His Gly Phe Asp Tyr Ala Lys Asp Tyr Phe Thr 180 185 190
- Asp Leu Ile Thr Asn Glu Ser Ile Asn Tyr Phe Lys Met Ser Lys Arg 195 200 205
- Met Tyr Pro His Arg Pro Val Met Met Val Ile Ser His Ala Ala Pro 210 215 220
- His Gly Pro Glu Asp Ser Ala Pro Gln Phe Ser Lys Leu Tyr Pro Asn 225 230 235 240
- Ala Ser Gln His Ile Thr Pro Ser Tyr Asn Tyr Ala Pro Asn Met Asp 245 250 255
- Lys His Trp Ile Met Gln Tyr Thr Gly Pro Met Leu Pro Ile His Met 260 265 270
- Glu Phe Thr Asn Ile Leu Gln Arg Lys Arg Leu Gln Thr Leu Met Ser 275 280 285
- Val Asp Asp Ser Val Glu Arg Leu Tyr Asn Met Leu Val Glu Thr Gly 290 295 300

- Glu Leu Glu Asn Thr Tyr Ile Ile Tyr Thr Ala Asp His Gly Tyr His 305 310 315 320
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- Asp Ile Arg Val Pro Phe Phe Ile Arg Gly Pro Ser Val Glu Pro Gly 340 345 350
- Ser Ile Val Pro Gln Ile Val Leu Asn Ile Asp Leu Ala Pro Thr Ile 355 360 365
- Leu Asp Ile Ala Gly Leu Asp Thr Pro Pro Asp Val Asp Gly Lys Ser 370 375 380
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- Lys Phe Leu Arg Lys Lys Glu Glu Ser Ser Lys Asn Ile Gln Gln Ser 420 425 430
- Asn His Leu Pro Lys Tyr Glu Arg Val Lys Glu Leu Cys Gln Gln Ala 435 440 445
- Arg Tyr Gln Thr Ala Cys Glu Gln Pro Gly Gln Lys Trp Gln Cys Ile 450 455 460
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- Asp Leu Leu Thr Val Arg Gln Ser Thr Arg Asn Leu Tyr Ala Arg Gly
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- Ala Ser Arg Ser Gln Arg Lys Ser Gln Arg Gln Phe Leu Arg Asn Gln
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- Gly Thr Pro Lys Tyr Lys Pro Arg Phe Val His Thr Arg Gln Thr Arg 530 535 540
- Ser Leu Ser Val Glu Phe Glu Gly Glu Ile Tyr Asp Ile Asn Leu Glu 545 550 555 560
- Glu Glu Glu Glu Leu Gln Val Leu Gln Pro Arg Asn Ile Ala Lys Arg 565 570 575

- His Asp Glu Gly His Lys Gly Pro Arg Asp Leu Gln Ala Ser Ser Gly 580 585 590
- Gly Asn Arg Gly Arg Met Leu Ala Asp Ser Ser Asn Ala Val Gly Pro 595 600 605
- Pro Thr Thr Val Arg Val Thr His Lys Cys Phe Ile Leu Pro Asn Asp 610 615 620
- Ser Ile His Cys Glu Arg Glu Leu Tyr Gln Ser Ala Arg Ala Trp Lys 625 630 635 640
- Asp His Lys Ala Tyr Ile Asp Lys Glu Ile Glu Ala Leu Gln Asp Lys 645 650 655
- Ile Lys Asn Leu Arg Glu Val Arg Gly His Leu Lys Arg Arg Lys Pro 660 665 670
- Glu Glu Cys Ser Cys Ser Lys Gln Ser Tyr Tyr Asn Lys Glu Lys Gly 675 680 685
- Val Lys Lys Gln Glu Lys Leu Lys Ser His Leu His Pro Phe Lys Glu 690 695 700
- Ala Ala Gln Glu Val Asp Ser Lys Leu Gln Leu Phe Lys Glu Asn Asn 705 710 715 720
- Arg Arg Arg Lys Lys Glu Arg Lys Glu Lys Arg Arg Gln Arg Lys Gly
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- Glu Glu Cys Ser Leu Pro Gly Leu Thr Cys Phe Thr His Asp Asn Asn 740 745 750
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- Phe Asp Met Asn Thr Asp Pro Tyr Gln Leu Thr Asn Thr Val His Thr 805 810 815
- Val Glu Arg Gly Ile Leu Asn Gln Leu His Val Gln Leu Met Glu Leu 820 825 830
- Arg Ser Cys Gln Gly Tyr Lys Gln Cys Asn Pro Arg Pro Lys Asn Leu 835 840 845

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<212> DNA

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<212> PRT
<213> Homo sapiens
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- <221> UNSURE
- <222> (111)
- <223> "Xaa" at position 111 can be any amino acid.

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Ser Cys Arg Ala Gly Thr Tyr Tyr Asp Gly Ala Arg Glu Arg Cys Ile

Leu Cys Pro Asn Gly Thr Phe Gln Asn Glu Glu Gly Gln Met Thr Cys 55

Glu Pro Cys Pro Arg Pro Gly Asn Ser Gly Ala Leu Lys Thr Pro Glu

- Ala Trp Asn Met Ser Glu Cys Gly Gly Leu Cys Gln Pro Thr Glu Tyr 85 90 95
- Ser Ala Asp Gly Phe Ala Pro Cys Gln Leu Cys Ala Leu Gly Xaa Phe 100 105 110
- Gln Pro Glu Ala Gly Arg Thr Ser Cys Phe Pro Cys Gly Gly Gly Leu 115 120 125
- Ala Thr Lys His Gln Gly Ala Thr Ser Phe Gln Asp Cys Glu Thr Arg 130 135 140
- Val Gln Cys Ser Pro Gly His Phe Tyr Asn Thr Thr Thr His Arg Cys 145 150 155 160
- Ile Arg Cys Pro Val Gly Thr Tyr Gln Pro Glu Phe Gly Lys Asn Asn 165 170 175
- Cys Val Ser Cys Pro Gly Asn Thr Thr Thr Asp Phe Asp Gly Ser Thr 180 185 190
- Asn Ile Thr Gln Cys Lys Asn Arg Arg Cys Gly Glu Leu Gly Asp 195 200 205
- Phe Thr Gly Tyr Ile Glu Ser Pro Asn Tyr Pro Gly Asn Tyr Pro Ala 210 215 220
- Asn Thr Glu Cys Thr Trp Thr Ile Asn Pro Pro Pro Lys Arg Arg Ile 225 230 235 240
- Leu Ile Val Val Pro Glu Ile Phe Leu Pro Ile Glu Asp Asp Cys Gly 245 250 255
- Asp Tyr Leu Val Met Arg Lys Thr Ser Ser Ser Asn Ser Val Thr Thr 260 265 270
- Tyr Glu Thr Cys Gln Thr Tyr Glu Arg Pro Ile Ala Phe Thr Ser Arg 275 280 285
- Ser Lys Lys Leu Trp Ile Gln Phe Lys Ser Asn Glu Gly Asn Ser Ala 290 295 300
- Arg Gly Phe Gln Val Pro Tyr Val Thr Tyr Asp Glu Asp Tyr Gln Glu 305 310 315 320
- Leu Ile Glu Asp Ile Val Arg Asp Gly Arg Leu Tyr Ala Ser Glu Asn 325 330 335
- His Gln Glu Ile Leu Lys Asp Lys Lys Leu Ile Lys Ala Leu Phe Asp 340 345 350

Val Leu Ala His Pro Gln Asn Tyr Phe Lys Tyr Thr Ala Gln Glu Ser 355 360 365

Arg Glu Met Phe Pro Arg Ser Phe Ile Arg Leu Leu Arg Ser Lys Val 370 375 380

Ser Arg Phe Leu Arg Pro Tyr Lys 385 390

<210> 39

<211> 392

<212> PRT

<213> Homo sapiens

<220>

<221> UNSURE

<222> (111)

<223> "Xaa" at position 111 can be any amino acid.

<400> 39

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Leu Cys Pro Asn Gly Thr Phe Gln Asn Glu Glu Gly Gln Met Thr Cys
50 55 60

Glu Pro Cys Pro Arg Pro Gly Asn Ser Gly Ala Leu Lys Thr Pro Glu 65 70 75 80

Ala Trp Asn Met Ser Glu Cys Gly Gly Leu Cys Gln Pro Thr Glu Tyr 85 90 95

Ser Ala Asp Gly Phe Ala Pro Cys Gln Leu Cys Ala Leu Gly Xaa Phe 100 105 110

Gln Pro Glu Ala Gly Arg Thr Ser Cys Phe Pro Cys Gly Gly Leu 115 120 125

Ala Thr Lys His Gln Gly Ala Thr Ser Phe Gln Asp Cys Glu Thr Arg 130 135 140

Val Gln Cys Ser Pro Gly His Phe Tyr Asn Thr Thr His Arg Cys 145 150 155 160 •

Ile Arg Cys Pro Val Gly Thr Tyr Gln Pro Glu Phe Gly Lys Asn Asn 165 170 175

Cys Val Ser Cys Pro Gly Asn Thr Thr Thr Asp Phe Asp Gly Ser Thr 180 185 190

Asn Ile Thr Gln Cys Lys Asn Arg Arg Cys Gly Glu Leu Gly Asp 195 200 205

Phe Thr Gly Tyr Ile Glu Ser Pro Asn Tyr Pro Gly Asn Tyr Pro Ala 210 215 220

Asn Thr Glu Cys Thr Trp Thr Ile Asn Pro Pro Pro Lys Arg Arg Ile 225 230 235 240

Leu Ile Val Val Pro Glu Ile Phe Leu Pro Ile Glu Asp Asp Cys Gly 245 250 255

Asp Tyr Leu Val Met Arg Lys Thr Ser Ser Ser Asn Ser Val Thr Thr 260 265 270

Tyr Glu Thr Cys Gln Thr Tyr Glu Arg Pro Ile Ala Phe Thr Ser Arg 275 280 285

Ser Lys Lys Leu Trp Ile Gln Phe Lys Ser Asn Glu Gly Asn Ser Ala 290 295 300

Arg Gly Phe Gln Val Pro Tyr Val Thr Tyr Asp Glu Asp Tyr Gln Glu 305 310 315 320

Leu Ile Glu Asp Ile Val Arg Asp Gly Arg Leu Tyr Ala Ser Glu Asn 325 330 335

His Gln Glu Ile Leu Lys Asp Lys Leu Ile Lys Ala Leu Phe Asp 340 345 350

Val Leu Ala His Pro Gln Asn Tyr Phe Lys Tyr Thr Ala Gln Glu Ser 355 360 365

Arg Glu Met Phe Pro Arg Ser Phe Ile Arg Leu Leu Arg Ser Lys Val 370 375 380

Ser Arg Phe Leu Arg Pro Tyr Lys 385 390

<210> 40

<211> 162

<212> PRT

<213> Mouse

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Ser Phe Ile Arg Leu Leu Arg Ser Lys Val Ser Arg Phe Leu Arg Pro

155

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150

Tyr Lys

145

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cggatgeea aceaeetgea ceaeagaeg caggagaaeg ceateetgge categageag 180
tacgaggage tggtggaegt gaaetgeage geegtgetge gettettett etgtgeeatg 240
tacgegeea tttgeaeeet ggagtteetg caegaeeeta teaageegtg caagteggtg 300
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<211> 346

<212> PRT

<213> Homo sapiens

<400> 43

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Arg His Met Pro Trp Asn Ile Thr Arg Met Pro Asn His Leu His His 35 40 45

Ser Thr Glu Asn Ala Ile Leu Ala Ile Glu Gln Tyr Glu Glu Leu 50 55 60

Val Asp Val Asn Cys Ser Ala Val Leu Arg Phe Phe Cys Ala Met 65 70 75 80

Tyr Ala Pro Ile Cys Thr Leu Glu Phe Leu His Asp Pro Ile Lys Pro 85 90 95

Cys Lys Ser Val Cys Gln Arg Ala Arg Asp Asp Cys Glu Pro Leu Met 100 105 110

Lys Met Tyr Asn His Ser Trp Pro Glu Ser Leu Ala Cys Asp Glu Leu 115 120 125

Pro Val Tyr Asp Arg Gly Val Cys Ile Ser Pro Glu Ala Ile Val Thr 130 135 140

Asp Leu Pro Glu Asp Val Lys Trp Ile Asp Ile Thr Pro Asp Met Met 145 150 155 160 Val Gln Glu Arg Pro Leu Asp Val Asp Cys Lys Arg Leu Ser Pro Asp 165 170 175

Arg Cys Lys Cys Lys Lys Val Lys Pro Thr Leu Ala Thr Tyr Leu Ser 180 185 190

Lys Asn Tyr Ser Tyr Val Ile His Ala Lys Ile Lys Ala Val Gln Arg 195 200 205

Ser Gly Cys Asn Glu Val Thr Thr Val Val Asp Val Lys Glu Ile Phe 210 215 220

Lys Ser Ser Ser Pro Ile Pro Arg Thr Gln Val Pro Leu Ile Thr Asn 225 230 235 240

Ser Ser Cys Gln Cys Pro His Ile Leu Pro His Gln Asp Val Leu Ile 245 250 255

Met Cys Tyr Glu Trp Arg Ser Arg Met Met Leu Leu Glu Asn Cys Leu 260 265 270

Val Glu Lys Trp Arg Asp Gln Leu Ser Lys Arg Ser Ile Gln Trp Glu 275 280 285

Glu Arg Leu Gln Glu Gln Arg Arg Thr Val Gln Asp Lys Lys Thr 290 295 300

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<212> DNA

<213> Homo sapiens

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749

501

Met Leu Asn Leu Met His Glu Thr Thr Asp Lys Asn Leu Ser Pro Asp

105

100

Gly Gln Tyr Val Pro Arg Ile Met Phe Val Asp Pro Ser Leu Thr Val 115 120 125

Arg Ala Asp Ile Ala Gly Arg Tyr Ser Asn Arg Leu Tyr Thr Tyr Glu 130 135 140

Pro Arg Asp Leu Pro Leu Leu Ile Glu Asn Met Lys Lys Ala Leu Arg 145 150 155 160

Leu Ile Gln Ser Glu Leu 165

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<213> Homo sapiens

<400> 48

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Arg Ser His Ala Ala Glu Ala Pro Gly Asp Pro Gln Ala Ala Ser 35 40 45

Leu Leu Ala Pro Met Asp Val Gly Glu Glu Pro Leu Glu Lys Ala Ala 50 55 60

Arg Ala Arg Thr Ala Lys Asp Pro Asn Thr Tyr Lys Val Leu Ser Leu 65 70 75 80

Val Leu Ser Val Cys Val Leu Thr Thr Ile Leu Gly Cys Ile Phe Gly

				85					90					95	
Leu	Lys	Pro	Ser 100	Cys	Ala	Lys	Glu	Val 105	Lys	Ser	Cys	Lys	Gly 110	Arg	Cys
Phe	Glu	Arg 115	Thr	Phe	Gly	Asn	Cys 120	Arg	Cys	Asp	Ala	Ala 125	Cys	Val	Glu
Leu	Gly 130	Asn	Cys	Cys	Leu	Asp 135	Tyr	Gln	Glu	Thr	Cys 140	Ile	Glu	Pro	Glu
His 145	Ile	Trp	Thr	Cys	Asn 150	Lys	Phe	Arg	Cys	Gly 155	Glu	Lys	Arg	Leu	Thr 160
Arg	Ser	Leu	Cys	Ala 165	Cys	Ser	Asp	Asp	Cys 170	Lys	Asp	Lys	Gly	Asp 175	Cys
Cys	Ile	Asn	Tyr 180	Ser	Ser	Val	Cys	Gln 185	Gly	Glu	Lys	Ser	Trp 190	Val	Glu
Glu	Pro	Cys 195	Glu	Ser	Ile	Asn	Glu 200	Pro	Gln	Cys	Pro	Ala 205	Gly	Phe	Glu
Thr	Pro 210	Pro	Thr	Leu	Leu	Phe 215	Ser	Leu	Asp	Gly	Phe 220	Arg	Ala	Glu	Tyr
Leu 225	His	Thr	Trp	Gly	Gly 230	Leu	Leu	Pro	Val	Ile 235	Ser	Lys	Leu	Lys	Lys 240
Cys	Gly	Thr	Tyr	Thr 245	Lys	Asn	Met	Arg	Pro 250	Val	Tyr	Pro	Thr	Lys 255	Thr
Phe	Pro	Asn	His 260	Tyr	Ser	Ile	Val	Thr 265	Gly	Leu	Tyr	Pro	Glu 270	Ser	His
Gly	Ile	Ile 275	Asp	Asn	Lys	Met	Tyr 280	Asp	Pro	Lys	Met	Asn 285	Ala	Ser	Phe
Ser	Leu 290	Lys	Ser	Lys	Glu	Lys 295	Phe	Asn	Pro	Glu	Trp 300	Tyr	Lys	Gly	Glu

340 345 350

Tyr Lys Met Tyr Asn Gly Ser Val Pro Phe Glu Glu Arg Ile Leu Ala

Pro Ile Trp Val Thr Ala Lys Tyr Gln Gly Leu Lys Ser Gly Thr Phe

Phe Trp Pro Gly Ser Asp Val Glu Ile Asn Gly Ile Phe Pro Asp Ile

310

325

305

Val Leu Gln Trp Leu Gln Leu Pro Lys Asp Glu Arg Pro His Phe Tyr

330

315

355 360 365

Thr Leu Tyr Leu Glu Glu Pro Asp Ser Ser Gly His Ser Tyr Gly Pro 370 380

Val Ser Ser Glu Val Ile Lys Ala Leu Gln Arg Val Asp Gly Met Val 385 390 395 400

Gly Met Leu Met Asp Gly Leu Lys Glu Leu Asn Leu His Arg Cys Leu 405 410 415

Asn Leu Ile Leu Ile Ser Asp His Gly Met Glu Gln Gly Ser Cys Lys 420 425 430

Lys Tyr Ile Tyr Leu Asn Lys Tyr Leu Gly Asp Val Lys Asn Ile Lys 435 440 445

Val Ile Tyr Gly Pro Ala Ala Arg Leu Arg Pro Ser Asp Val Pro Asp 450 455 460

Lys Tyr Tyr Ser Phe Asn Tyr Glu Gly Ile Ala Arg Asn Leu Ser Cys 465 470 475 480

Arg Glu Pro Asn Gln His Phe Lys Pro Tyr Leu Lys His Phe Leu Pro
485 490 495

Lys Arg Leu His Phe Ala Lys Ser Asp Arg Ile Glu Pro Leu Thr Phe 500 505 510

Tyr Leu Asp Pro Gln Trp Gln Leu Ala Leu Asn Pro Ser Glu Arg Lys 515 520 525

Tyr Cys Gly Ser Gly Phe His Gly Ser Asp Asn Val Phe Ser Asn Met 530 540

Gln Ala Leu Phe Val Gly Tyr Gly Pro Gly Phe Lys His Gly Ile Glu 545 550 555 560

Ala Asp Thr Phe Glu Asn Ile Glu Val Tyr Asn Leu Met Cys Asp Leu 565 570 575

Leu Asn Leu Thr Pro Ala Pro Asn Asn Gly Thr His Gly Ser Leu Asn 580 585 590

His Leu Leu Lys Asn Pro Val Tyr Thr Pro Lys His Pro Lys Glu Val 595 600 605

His Pro Leu Val Gln Cys Pro Phe Thr Arg Asn Pro Arg Asp Asn Leu 610 615 620

Gly Cys Ser Cys Asn Pro Ser Ile Leu Pro Ile Glu Asp Phe Gln Thr

625 630 635 640

Gln Phe Asn Leu Thr Val Ala Glu Glu Lys Ile Ile Lys His Glu Thr 645 650 655

Leu Pro Tyr Gly Arg Pro Arg Val Leu Gln Lys Glu Asn Thr Ile Cys
660 665 670

Leu Leu Ser Gln His Gln Phe Met Ser Gly Tyr Ser Gln Asp Ile Leu 675 680 685

Met Pro Leu Trp Thr Ser Tyr Thr Val Asp Arg Asn Asp Ser Phe Ser 690 695 700

Thr Glu Asp Phe Ser Asn Cys Leu Tyr Gln Asp Phe Arg Ile Pro Leu 705 710 715 720

Ser Pro Val His Lys Cys Ser Phe Tyr Lys Asn Asn Thr Lys Val Ser 725 730 735

Tyr Gly Phe Leu Ser Pro Pro Gln Leu Asn Lys Asn Ser Ser Gly Ile 740 745 750

Tyr Ser Glu Ala Leu Leu Thr Thr Asn Ile Val Pro Met Tyr Gln Ser 755 760 765

Phe Gln Val Ile Trp Arg Tyr Phe His Asp Thr Leu Leu Arg Lys Tyr 770 775 780

Ala Glu Glu Arg Asn Gly Val Asn Val Val Ser Gly Pro Val Phe Asp 785 790 795 800

Phe Asp Tyr Asp Gly Arg Cys Asp Ser Leu Glu Asn Leu Arg Gln Lys 805 810 815

Arg Arg Val Ile Arg Asn Gln Glu Ile Leu Ile Pro Thr His Phe 820 825 830

Ile Val Leu Thr Ser Cys Lys Asp Thr Ser Gln Thr Pro Leu His Cys 835 840 845

Glu Asn Leu Asp Thr Leu Ala Phe Ile Leu Pro His Arg Thr Asp Asn 850 855 860

Ser Glu Ser Cys Val His Gly Lys His Asp Ser Ser Trp Val Glu Glu 865 870 875 880

Leu Leu Met Leu His Arg Ala Arg Ile Thr Asp Val Glu His Ile Thr 885 890 895

Gly Leu Ser Phe Tyr Gln Gln Arg Lys Glu Pro Val Ser Asp Ile Leu

900 905 910

Lys Leu Lys Thr His Leu Pro Thr Phe Ser Gln Glu Asp 915 920 925

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<211> 299

<212> PRT

<213> Homo sapiens

<400> 50

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20 25 30

Ile Pro Thr Phe Ile Cys Ser Val Ile Tyr Leu Leu Ile Val Trp Leu
35 40 45

Gly Pro Lys Tyr Met Arg Asn Lys Gln Pro Phe Ser Cys Arg Gly Ile 50 55 60

Leu Val Val Tyr Asn Leu Gly Leu Thr Leu Leu Ser Leu Tyr Met Phe 65 70 75 80

Cys Glu Leu Val Thr Gly Val Trp Glu Gly Lys Tyr Asn Phe Phe Cys 85 90 95

Gln Gly Thr Arg Thr Ala Gly Glu Ser Asp Met Lys Ile Ile Arg Val 100 105 110

Leu Trp Trp Tyr Tyr Phe Ser Lys Leu Ile Glu Phe Met Asp Thr Phe 115 120 125

Phe Phe Ile Leu Arg Lys Asn Asn His Gln Ile Thr Val Leu His Val 130 135 140

Tyr His His Ala Ser Met Leu Asn Ile Trp Trp Phe Val Met Asn Trp 145 150 155 160

Val Pro Cys Gly His Ser Tyr Phe Gly Ala Thr Leu Asn Ser Phe Ile 165 170 175

His Val Leu Met Tyr Ser Tyr Tyr Gly Leu Ser Ser Val Pro Ser Met 180 185 190 Arg Pro Tyr Leu Trp Trp Lys Lys Tyr Ile Thr Gln Gly Gln Leu Leu 195 200 205

Gln Phe Val Leu Thr Ile Ile Gln Thr Ser Cys Gly Val Ile Trp Pro 210 215 220

Cys Thr Phe Pro Leu Gly Trp Leu Tyr Phe Gln Ile Gly Tyr Met Ile 225 230 235 240

Ser Leu Ile Ala Leu Phe Thr Asn Phe Tyr Ile Gln Thr Tyr Asn Lys 245 250 255

Lys Gly Ala Ser Arg Arg Lys Asp His Leu Lys Asp His Gln Asn Gly 260 265 270

Ser Met Ala Ala Val Asn Gly His Thr Asn Ser Phe Ser Pro Leu Glu 275 280 285

Asn Asn Val Lys Pro Arg Lys Leu Arg Lys Asp 290 295

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<211> 1019

<212> DNA

<213> Homo sapiens

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<211> 1332

<212> DNA

<213> Homo sapiens

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caaqttqaaa tacctttctt gtgttttcta attagaaaag taatatctac tcattgtaaa 180
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      species.
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Trp Ser Xaa Trp Ser
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												ccc Pro	794
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												aac Asn 250	890
												aca Thr	938
												aac Asn	986
	_		-									ctg Leu	1034
												tca Ser	1082
												agt Ser 330	1130
												gtg Val	1178

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agt ggt gat gct ttt tta cac ctt ctt cca cat tct cat gca agt cac

Ser	Gly 365	Asp	Ala	Phe	Leu	His 370	Leu	Leu	Pro	His	Ser 375	His	Ala	Ser	His	
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														gcc Ala 410		1370
														tat Tyr		1418
														aaa Lys		1466
														gtg Val		1514
														aat Asn		1562
														gca Ala 490		1610
														ttg Leu		1658
														tac Tyr		1706
														ttc Phe		1754
				_		_	_							gac Asp		1802
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His Ser Gln Arg Tyr Ser Arg Glu Glu Leu Lys Asp Ala Gly Val A 575 580 585	la
act ttg gcc tgg atg gtg ata atg ggt gat ggc ctg cac aat ttc ag Thr Leu Ala Trp Met Val Ile Met Gly Asp Gly Leu His Asn Phe Se 590 595 600	
gat ggc cta gca att ggt gct gct ttt act gaa ggc tta tca agt gg Asp Gly Leu Ala Ile Gly Ala Ala Phe Thr Glu Gly Leu Ser Ser G 605 610 615	
tta agt act tct gtt gct gtg ttc tgt cat gag ttg cct cat gaa tt Leu Ser Thr Ser Val Ala Val Phe Cys His Glu Leu Pro His Glu Le 620 625 630 63	
ggt gac ttt gct gtt cta cta aag gct ggc atg acc gtt aag cag gc Gly Asp Phe Ala Val Leu Leu Lys Ala Gly Met Thr Val Lys Gln Al 640 645 650	
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ttt gca ctt act gct ggc tta ttc atg tat gtt gct ctg gtt gat at Phe Ala Leu Thr Ala Gly Leu Phe Met Tyr Val Ala Leu Val Asp Me 685 690 695	_
gta cct gaa atg ctg cac aat gat gct agt gac cat gga tgt agc cg Val Pro Glu Met Leu His Asn Asp Ala Ser Asp His Gly Cys Ser Ar 700 705 710 72	•
tgg ggg tat ttc ttt tta cag aat gct ggg atg ctt ttg ggt ttt gg Trp Gly Tyr Phe Phe Leu Gln Asn Ala Gly Met Leu Leu Gly Phe Gl 720 725 730	
att atg tta ctt att tcc ata ttt gaa cat aaa atc gtg ttt cgt at Ile Met Leu Leu Ile Ser Ile Phe Glu His Lys Ile Val Phe Arg II 735 740 745	
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Gln Val Pro Leu Asn Ala Thr Glu Phe Asn Tyr Leu Cys Pro Ala Ile 280 285 275

Ile Asn Gln Ile Asp Ala Arg Ser Cys Leu Ile His Thr Ser Glu Lys 295 300 Lys Ala Glu Ile Pro Pro Lys Thr Tyr Ser Leu Gln Ile Ala Trp Val 310 315 Gly Gly Phe Ile Ala Ile Ser Ile Ile Ser Phe Leu Ser Leu Leu Gly 325 330 Val Ile Leu Val Pro Leu Met Asn Arg Val Phe Phe Lys Phe Leu Leu 340 345 Ser Phe Leu Val Ala Leu Ala Val Gly Thr Leu Ser Gly Asp Ala Phe 360 Leu His Leu Leu Pro His Ser His Ala Ser His His Ser His Ser 375 380 His Glu Glu Pro Ala Met Glu Met Lys Arg Gly Pro Leu Phe Ser His 390 395 Leu Ser Ser Gln Asn Ile Glu Glu Ser Ala Tyr Phe Asp Ser Thr Trp 405 410 Lys Gly Leu Thr Ala Leu Gly Gly Leu Tyr Phe Met Phe Leu Val Glu His Val Leu Thr Leu Ile Lys Gln Phe Lys Asp Lys Lys Lys Asn 440 Gln Lys Lys Pro Glu Asn Asp Asp Val Glu Ile Lys Lys Gln Leu 455 Ser Lys Tyr Glu Ser Gln Leu Ser Thr Asn Glu Glu Lys Val Asp Thr 470 475 Asp Asp Arg Thr Glu Gly Tyr Leu Arg Ala Asp Ser Gln Glu Pro Ser 490 His Phe Asp Ser Gln Gln Pro Ala Val Leu Glu Glu Glu Val Met 500 505 Ile Ala His Ala His Pro Gln Glu Val Tyr Asn Glu Tyr Val Pro Arg 520 Gly Cys Lys Asn Lys Cys His Ser His Phe His Asp Thr Leu Gly Gln 535 Ser Asp Asp Leu Ile His His His Asp Tyr His His Ile Leu His 550 His His His Gln Asn His His Pro His Ser His Ser Gln Arg Tyr 565 570 Ser Arg Glu Glu Leu Lys Asp Ala Gly Val Ala Thr Leu Ala Trp Met 585 Val Ile Met Gly Asp Gly Leu His Asn Phe Ser Asp Gly Leu Ala Ile 600 Gly Ala Ala Phe Thr Glu Gly Leu Ser Ser Gly Leu Ser Thr Ser Val 615 Ala Val Phe Cys His Glu Leu Pro His Glu Leu Gly Asp Phe Ala Val 630 635 Leu Leu Lys Ala Gly Met Thr Val Lys Gln Ala Val Leu Tyr Asn Ala 650 Leu Ser Ala Met Leu Ala Tyr Leu Gly Met Ala Thr Gly Ile Phe Ile Gly His Tyr Ala Glu Asn Val Ser Met Trp Ile Phe Ala Leu Thr Ala Gly Leu Phe Met Tyr Val Ala Leu Val Asp Met Val Pro Glu Met Leu

700 690 695 His Asn Asp Ala Ser Asp His Gly Cys Ser Arg Trp Gly Tyr Phe Phe 710 715 Leu Gln Asn Ala Gly Met Leu Leu Gly Phe Gly Ile Met Leu Leu Ile 730 725 Ser Ile Phe Glu His Lys Ile Val Phe Arg Ile Asn Phe 745 740 <210> 56 <211> 293 <212> DNA <213> Homo sapiens <220> <221> misc feature <222> (190) <223> "n" at position 190 can be any base <400> 56 tttttttac tgtgaaattc taaaatcata tttattcacc aattcacaga aagtgtcata 60 acgaccacca acatgaatca gtttgtaggc atttacaagc cacagctgaa aataaaaatc 120 tgtctgtgtt gaataggcat ttaacaaatt acttgaaaac tgcaagaatc ataattatta 180 taaatttaan gtttgtgatt caaacatggg taagatcaca gtcatgggga gaagcccaac 240 agattcctgt gatgatcctt attttcttcc tatatcttta tatacatagg agg <210> 57 <211> 2053 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (85)..(1347) <400> 57 ceggetegeg ceeteeggge ceageeteec gageettegg agegggegee gteecageec 60 ageteegggg aaaegegage egeg atg eet ggg ggg tge tee egg gge eee 111 Met Pro Gly Gly Cys Ser Arg Gly Pro gee gee ggg gae ggg egt etg egg etg geg ega eta geg etg gta ete 159 Ala Ala Gly Asp Gly Arg Leu Arg Leu Ala Arg Leu Ala Leu Val Leu 10

35

207

40

ctg ggc tgg gtc tcc tcg tct tct ccc acc tcc tcg gca tcc tcc ttc

Leu Gly Trp Val Ser Ser Ser Pro Thr Ser Ser Ala Ser Ser Phe

30

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						gcg Ala									303
	-	_	_			cgc Arg 80									351
						ctc Leu									399
		_		_		gcc Ala	_			_					447
				_		agc Ser	-	_	_		_	_			495
			_		_	ctg Leu	_	_		_		_			543
						ttc Phe 160									591
_	_		_			gtg Val									639
						aac Asn									687
						gca Ala									735
						tac Tyr									783
	_					gac Asp 240									831

acc tac gtg to Thr Tyr Val Se 250						879
gag gac aat go Glu Asp Asn Al						927
caa ggt cta co Gln Gly Leu Pr 28	o His Ile	Arg Val Ph				975
tgc gac tgc ca Cys Asp Cys Hi 300		-				1023
gta gtg cag gg Val Val Gln Gl 315						1071
agg aat cgg gt Arg Asn Arg Va 330						1119
ccg att ctt cc Pro Ile Leu Pr						1167
gtt tta gcc ct Val Leu Ala Le 36	u Ile Gly	Ala Ile Ph				1215
cgc aag ggg at Arg Lys Gly Il 380						1263
gat cac atg ga Asp His Met Gl 395						1311
aga tta aca aa Arg Leu Thr As 410	-			tga gaaatat	tag	1357
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<211> 420

<212> PRT

<213> Homo sapiens

<400> 58

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Leu Pro Arg Asp Val Leu Ala Gln Leu Pro Ser Leu Arg His Leu Asp 225 230 235 Leu Ser Asn Asn Ser Leu Val Ser Leu Thr Tyr Val Ser Phe Arg Asn 245 250 Leu Thr His Leu Glu Ser Leu His Leu Glu Asp Asn Ala Leu Lys Val 265 Leu His Asn Gly Thr Leu Ala Glu Leu Gln Gly Leu Pro His Ile Arg 280 285 Val Phe Leu Asp Asn Asn Pro Trp Val Cys Asp Cys His Met Ala Asp 300 295 Met Val Thr Trp Leu Lys Glu Thr Glu Val Val Gln Gly Lys Asp Arg 310 315 Leu Thr Cys Ala Tyr Pro Glu Lys Met Arg Asn Arg Val Leu Leu Glu 325 330 Leu Asn Ser Ala Asp Leu Asp Cys Asp Pro Ile Leu Pro Pro Ser Leu 345 Gln Thr Ser Tyr Val Phe Leu Gly Ile Val Leu Ala Leu Ile Gly Ala 360 Ile Phe Leu Leu Val Leu Tyr Leu Asn Arg Lys Gly Ile Lys Lys Trp 375 380 Met His Asn Ile Arg Asp Ala Cys Arg Asp His Met Glu Gly Tyr His 390 395 Tyr Arg Tyr Glu Ile Asn Ala Asp Pro Arg Leu Thr Asn Leu Ser Ser 410 Asn Ser Asp Val 420

420

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<211> 232

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<213> Homo sapiens

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<211> 281

<212> DNA

<213> Homo sapiens

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281

627

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Gly Gln Leu Glu Gln Asn Val Tyr Gly Ile Thr Ile Ile Ser Gly Leu

110 115 120

						aat Asn										675
						gat Asp										723
						gaa Glu										771
						ccc Pro										819
						gaa Glu 195										867
	_	_				cat His									_	915
Lys	Pro	Asn	Pro	Pro 225	His	aat Asn	Leu	Ser	Val 230	Ile	Asn	Ser	Glu	Glu 235	Leu	963
						aca Thr										1011
						att Ile							_		_	1059
						gaa Glu 275										1107
						cct Pro										1155
						aag Lys										1203
gca	agt	aaa	atc	acc	tat	gaa	gat	aga	cca	tct	aaa	gca	cca	agt	ttc	1251

Ala	Ser	Gly	Ile 320	Thr	Tyr	Glu	Asp	Arg 325	Pro	Ser	Lys	Ala	Pro 330	Ser	Phe	
		aaa Lys 335														1299
		tgg Trp														1347
_		gaa Glu					_									1395
		aat Asn														1443
	_	acc Thr			_	-										1491
		act Thr 415														1539
_		aaa Lys	_				-									1587
		agg Arg	_		_	_										1635
		aaa Lys														1683
		cgc Arg														1731
_		aca Thr 495	_													1779
		aag Lys														1827

_						Gly 333									1875
			~	_	_	cag Gln					_				1923
		_				gga Gly		_		_			 -		1971
			_			ttg Leu			_						2019
						tac Tyr 595									2067
_						cca Pro	_		_			_	_	-	2115
	_			-	_	tta Leu	_			_				_	2163
	_					aag Lys							_		2211
		_		_		tca Ser	_	_			_	_			2259
						aat Asn 675									2307
_					_	gta Val	_	_	_						2355
	_				_	gat Asp	_			_	_	_		_	2403
						gga Gly									2451

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		Pro Ser Val	caa gtc ttc tca aga tcc Gln Val Phe Ser Arg Ser 775 780	2595
			gag cgg cca gaa gat cta Glu Arg Pro Glu Asp Leu 795	2643
	o His Val Asp		ggt att ttg ccc agg caa Gly Ile Leu Pro Arg Gln 810	2691
			gaa tcc agt cca gat att Glu Ser Ser Pro Asp Ile 825	2739
			tca gtc aat gag gaa gat Ser Val Asn Glu Glu Asp 840	2787
		Ile Ser Asp	cat att tca caa tcc tgt His Ile Ser Gln Ser Cys 855 860	2835
			gtt tct gca gca gat gct Val Ser Ala Ala Asp Ala 875	2883
	y Thr Glu Gly		aga ttt gaa aca gtt ggc Arg Phe Glu Thr Val Gly 890	2931
			aaa agt tac tta cca cag Lys Ser Tyr Leu Pro Gln 905	2979
act gta cgg ca Thr Val Arg Gl 910		Met Pro Gln	tga aggactagta gttcctgcta	3032
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<210> 62

330

Asp Pro Ser His Thr Gln Gly Tyr Arg Thr Val Gln Leu Val Trp Lys 340 345 350

Thr Leu Pro Pro Phe Glu Ala Asn Gly Lys Ile Leu Asp Tyr Glu Val

325

Thr Leu Thr Arg Trp Lys Ser His Leu Gln Asn Tyr Thr Val Asn Ala Thr Lys Leu Thr Val Asn Leu Thr Asn Asp Arg Tyr Leu Ala Thr Leu Thr Val Arq Asn Leu Val Gly Lys Ser Asp Ala Ala Val Leu Thr Ile Pro Ala Cys Asp Phe Gln Ala Thr His Pro Val Met Asp Leu Lys Ala . Phe Pro Lys Asp Asn Met Leu Trp Val Glu Trp Thr Thr Pro Arg Glu Ser Val Lys Lys Tyr Ile Leu Glu Trp Cys Val Leu Ser Asp Lys Ala Pro Cys Ile Thr Asp Trp Gln Glu Asp Gly Thr Val His Arg Thr Tyr Leu Arg Gly Asn Leu Ala Glu Ser Lys Cys Tyr Leu Ile Thr Val Thr Pro Val Tyr Ala Asp Gly Pro Gly Ser Pro Glu Ser Ile Lys Ala Tyr Leu Lys Gln Ala Pro Pro Ser Lys Gly Pro Thr Val Arg Thr Lys Lys Val Gly Lys Asn Glu Ala Val Leu Glu Trp Asp Gln Leu Pro Val Asp Val Gln Asn Gly Phe Ile Arg Asn Tyr Thr Ile Phe Tyr Arg Thr Ile Ile Gly Asn Glu Thr Ala Val Asn Val Asp Ser Ser His Thr Glu Tyr Thr Leu Ser Ser Leu Thr Ser Asp Thr Leu Tyr Met Val Arg Met Ala Ala Tyr Thr Asp Glu Gly Gly Lys Asp Gly Pro Glu Phe Thr Phe Thr Thr Pro Lys Phe Ala Gln Gly Glu Ile Glu Ala Ile Val Val Pro Val Cys Leu Ala Phe Leu Leu Thr Thr Leu Leu Gly Val Leu Phe Cys - 630 Phe Asn Lys Arg Asp Leu Ile Lys Lys His Ile Trp Pro Asn Val Pro Asp Pro Ser Lys Ser His Ile Ala Gln Trp Ser Pro His Thr Pro Pro Arg His Asn Phe Asn Ser Lys Asp Gln Met Tyr Ser Asp Gly Asn Phe Thr Asp Val Ser Val Val Glu Ile Glu Ala Asn Asp Lys Lys Pro Phe Pro Glu Asp Leu Lys Ser Leu Asp Leu Phe Lys Lys Glu Lys Ile Asn Thr Glu Gly His Ser Ser Gly Ile Gly Gly Ser Ser Cys Met Ser Ser Ser Arg Pro Ser Ile Ser Ser Ser Asp Glu Asn Glu Ser Ser Gln Asn Thr Ser Ser Thr Val Gln Tyr Ser Thr Val Val His Ser Gly Tyr Arg 

His Gln Val Pro Ser Val Gln Val Phe Ser Arg Ser Glu Ser Thr Gln Pro Leu Leu Asp Ser Glu Glu Arg Pro Glu Asp Leu Gln Leu Val Asp His Val Asp Gly Gly Asp Gly Ile Leu Pro Arg Gln Gln Tyr Phe Lys Gln Asn Cys Ser Gln His Glu Ser Ser Pro Asp Ile Ser His Phe Glu Arg Ser Lys Gln Val Ser Ser Val Asn Glu Glu Asp Phe Val Arg Leu Lys Gln Gln Ile Ser Asp His Ile Ser Gln Ser Cys Gly Ser Gly Gln Met Lys Met Phe Gln Glu Val Ser Ala Ala Asp Ala Phe Gly Pro Gly Thr Glu Gly Gln Val Glu Arg Phe Glu Thr Val Gly Met Glu Ala Ala Thr Asp Glu Gly Met Pro Lys Ser Tyr Leu Pro Gln Thr Val Arg Gln Gly Gly Tyr Met Pro Gln